



Clean Version of Amended Claims
Pursuant to 37 C.F.R. § 1.121(c)(1)(ii)

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1. (Amended) A method for treating cells or biological materials to alter permeability, cell viability or structural integrity comprising

(a) administering acoustic energy to the biological materials at one or more frequencies;

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(b) measuring a property or the effect of the acoustic energy during the treatment with acoustic energy; and

(c) using the measurement obtained in step (b) to modify continued or subsequent application of acoustic energy to the biological materials during the treatment as needed to enhance the treatment.

2. (amended) The method of claim 1 wherein the property of the acoustic energy being measured in step b is one or more properties selected from the group consisting of pressure at one or more frequencies, and energy input at one or more frequencies.

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3. (amended) The method of claim 1 wherein the acoustic energy is effective to alter permeability of the biological materials to a chemical or biological agent selected from the group consisting of peptides, proteins, sugars, polysaccharides, nucleotides, polynucleotide molecules, synthetic organic compounds, synthetic inorganic compounds, endogenous organic compounds, endogenous inorganic compounds and combinations and aggregates thereof.

4. The method of claim 3 wherein the agent is in a form selected from the group consisting of cells or virus particles, nano or microparticles, liposomes or other lipid vesicles or emulsions.

sub c3 5. The method of claim 3 wherein the chemical or biological agent is delivered to cells or tissue.

B1 6. The method of claim 3 wherein the chemical or biological agent is detected or quantitated, further comprising
removing biological fluid or molecules simultaneously, previously, or subsequently to the application of acoustic energy, and
assaying the biological fluid or molecules to detect or quantitate the chemical or biological agents.

7. The method of claim 1 wherein the acoustic energy is administered to kill cells.

sub c4 8. The method of claim 1 wherein the biological materials are made more permeable by the exposure to acoustic energy.

9. (amended) The method of claim 8 wherein the biological materials are made partially or completely reversibly permeable.

10. The method of claim 1 wherein the biological materials are biological membranes.

11. The method of claim 1 wherein the biological material is skin.

12. The method of claim 1 wherein the acoustic energy is applied to biological materials in an amount effective to disaggregate or dissociate the materials.

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13. The method of claim 1 wherein the biological materials are blood vessels.

14. The method of claim 1 wherein the acoustic energy is applied at a frequency between 1 kHz and 10 MHz.

15. The method of claim 1 wherein the acoustic energy is ultrasound.

16. The method of claim 1 wherein the acoustic energy is applied at a peak positive pressure of up to 100 atmospheres.

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C5 17. (amended) The method of claim 1 wherein the acoustic energy is applied under conditions to effect cavitation within or on the surface of the biological materials.

B1 18. The method of claim 1 further comprising administering an agent to enhance transport within or permeability of the biological materials.

19. (Amended) The method of claim 1 wherein the property of the acoustic energy that is measured is measured at one or more frequencies other than the frequency or frequencies at which the acoustic energy is applied.

20. (Amended) The method of claim 1 wherein the property of the acoustic energy that is measured is measured at a frequency or frequencies corresponding to integer multiples of one-half or one-fourth of the frequency applied.

21. (Amended) The method of claim 1 wherein the acoustic energy is measured at one or more frequencies in the acoustic spectrum which do not correspond to peaks in the acoustic spectrum and are taken from the broadband signal of the acoustic spectrum.

22. (Amended) The method of claim 19 wherein the acoustic energy measurement is analyzed using a mathematical algorithm, selected from the group consisting of Fourier Transform and Fast Fourier Transform.

23. The method of claim 1 wherein the application of the acoustic energy is modified by changing an acoustic parameter selected from the group consisting of pressure, energy, frequency, pulse length, total exposure time, duty cycle, and combinations thereof.

24. The method of claim 1 wherein the application of the acoustic energy is modified by changing a non-acoustic parameter selected from the group consisting of temperature, fluid gas content, administration rate of molecules to be transported, sample collection rate, device position, and combinations thereof.

B1 25. The method of claim 1 wherein the application of the acoustic energy input is modified by interrupting the application.

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C6 26. (Twice Amended) A device comprising
(a) means for treating cells or biological materials by administering acoustic energy to the cells or biological materials to alter permeability, cell viability or structural integrity of biological materials at one or more frequencies;
(b) means for measuring a property or the effect of the acoustic energy during the treatment with acoustic energy; and
(c) means for using the measurement of the property of the acoustic energy to modify continued or subsequent application of acoustic energy to the biological materials during the treatment as needed to enhance the treatment.

c6 27. (Twice Amended) A method for altering cell viability or transport of chemical or biological agents into or through an internal organ, internal tissue or vessel in a human or other animal using acoustic energy, comprising:

administering acoustic energy at one or more frequencies by applying a transducer to a site on the human or other animal other than where transport or cell viability is to be altered; wherein the acoustic energy is effective to alter transport or cell viability at the internal organ, tissue or vessel.

B1 28. (Amended) The method of claim 27 wherein the acoustic energy is applied to the skin or a mucosal membrane and alters transport or cell viability at an internal organ, tissue or vessel.

29. (Amended) The method of claim 27 wherein the acoustic energy alters transport or cell viability of tumor cells.

30. (Amended) The method of claim 27 wherein the acoustic energy alters transport into or out of the cells of molecules selected from the group consisting of therapeutic, prophylactic and diagnostic agents.

sub 31 31. (Amended) The method of claim 27 wherein the transducer is placed inside the body using invasive or minimally invasive means.

Please add the following new claims 32 and 33.

sub 32 32. The method of claim 31 wherein the transducer is placed within a blood vessel using a catheter.

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33. The method of claim 31 wherein the transducer is placed within a surgical
incision.
